

## RAW SEQUENCE LISTING ERROR REPORT

The Biotechnology Systems Branch of the Scientific and Technical Information Center (STIC) detected errors when processing the following computer readable form:

Application Serial Number:	10/009.500
Source:	1/4/10
Date Processed by STIC:	6/6/2002.

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FOR CRF SUBMISSION QUESTIONS, PLEASE CONTACT MARK SPENCER, 703-308-4212.

FOR SEQUENCE RULES INTERPRETATION, PLEASE CONTACT ROBERT WAX, 703-308-4216. PATENTIN 2.1 e-mail help: patin21help@uspto.gov or phone 703-306-4119 (R. Wax) PATENTIN 3.0 e-mail help: patin3help@uspto.gov or phone 703-306-4119 (R. Wax)

TO REDUCE ERRORED SEQUENCE LISTINGS, PLEASE USE THE CHECKER VERSION 3.1 PROGRAM, ACCESSIBLE THROUGH THE U.S. PATENT AND TRADEMARK OFFICE WEBSITE. SEE BELOW FOR ADDRESS:

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Any reply including a sequence listing in electronic form should NOT be sent to the 20231 zip code address for the United States Patent and Trademark Office, and instead should be sent via the following to the indicated addresses:

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U.S. Patent and Trademark Office, Technology Center 1600, Reception Area, 7<sup>th</sup> Floor, Examiner Name, Sequence Information, Crystal Mall One, 1911 South Clark Street, Arlington, VA 22202

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Revised 01/29/2002



PCT10

RAW SEQUENCE LISTING

DATE: 06/06/2002

PATENT APPLICATION: US/10/009,500

TIME: 14:28:17

Input Set : A:\merck 2332 seq list

Output Set: N:\CRF3\06062002\J009500.raw

Does Not Lomply Corrected Diskette Needer /

5 <120> TITLE OF INVENTION: Hyaluronidase from the Hirudinaria manillensis, isolation, purification and recombinant method of production

9 <130> FILE REFERENCE: Manillase

C√-> 11 <140> CURRENT APPLICATION NUMBER: US/10/009,500

SA-> 12 <141> CURRENT FILING DATE: 2002-04-08

14 <160> NUMBER OF SEQ ID NOS: 15 16 <170> SOFTWARE: PatentIn Ver. 2.1

## ERRORED SEQUENCES

258 <210> SEQ ID NO: 3 259 <211> LENGTH: 488 260 <212> TYPE: PRT 261 <213> ORGANISM: Leech 263 <400> SEQUENCE: 3 264 Lys Glu Ile Ala Val Thr Ile Asp Asp Lys Asn Val Ile Ala Ser Ala 265 1 5 10 267 Ser Gly Ser Phe Leu Gly Val Ala Phe Asp Ala Ser Leu Phe Ser Pro 20 25 270 Lys Gly Leu Trp Ser Phe Val Asp Ile Thr Ser Pro Lys Leu Phe Lys

274 55 276 Ala Asn Trp Leu Phe Phe Asp Leu Asp Glu Asn Asn Lys Trp Lys Asp 277 65 70 75 279 Tyr Trp Ala Phe Lys Asp Lys Thr Pro Glu Thr Ala Thr Ile Thr Arg

40 273 Leu Leu Glu Gly Leu Ser Pro Gly Tyr Phe Arg Val Gly Gly Thr Phe

280 85 90 282 Arg Trp Leu Phe Arg Lys Gln Asn Asn Leu Lys Lys Glu Thr Phe Asp 100 105

185 Asn Leu Val Lys Leu Thr Lys Gly Ser Lys Met Arg Leu Leu Phe Asp 120

288 Leu Asn Ala Glu Val Arg Thr Gly Tyr Glu Ile Gly Lys Lys Met Thr 135 140 289 130

291 Ser Thr Trp Asp Ser Ser Glu Ala Glu Lys Leu Phe Lys Tyr Cys Val 150 155 160

294 Ser Lys Gly Tyr Gly Asp Asn Ile Asp Trp Glu Leu Gly Asn Glu Pro 295 165 170 175

297 Asp His Thr Ser Ala His Asn Leu Thr Glu Lys Gln Val Gly Glu Asp 180 185

300 Phe Lys Ala Leu His Lys Val Leu Glu Lys Tyr Pro Thr Leu Asn Lys

RAW SEQUENCE LISTING

PATENT APPLICATION: US/10/009,500

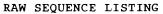
DATE: 06/06/2002 TIME: 14:28:17

Input Set : A:\merck 2332 seq list

Output Set: N:\CRF3\06062002\J009500.raw

195 200 301 E--> 303 Gly Ser Leu Val Gly Pro Asp Val Gly Trp Met Gly Val Ser Xaa Val 220. 210 225 E--> 306 Lys Gly Leu Ala Asp Glu Xaa Gly Asp His Val (Xaa Ala Phe Thr Leu 230 235 E--> 309 His Gln Tyr Tyr Phe Asp Gly Asn Xaa Ser Asp Val Ser Ile Tyr Leu 250 245 312 Asp Ala Thr Tyr Phe Lys Lys Leu Gln Gln Leu Phe Asp Lys Val Lys 260 265 315 Asp Val Leu Lys Asp Ser Pro His Lys Asp Glu Pro Leu Trp Leu Gly 316 275 280 318 Glu Thr Ser Ser Gly Tyr Asn Ser Gly Thr Glu Asp Val Ser Asp Arg 295 321 Tyr Val Ser Gly Phe Leu Thr Leu Asp Lys Leu Gly Leu Ser Ala Ala 310 324 Asn Asn Val Lys Val Val Ile Arg Gln Thr Ile Tyr Asn Gly Tyr Tyr 325 330 327 Gly Leu Leu Asp Lys Asn Thr Leu Glu Pro Asn Pro Asp Tyr Trp Leu 340 345 330 Met His Val His Asn Ser Leu Val Gly Asn Thr Val Phe Lys Val Asp 355 331 360 333 Val Ser Asp Pro Thr Asn Lys Ala Arg Val Tyr Ala Gln Cys Thr Lys 370 375 380 336 Thr Asn Ser Lys His Thr Gln Ser Arg Tyr Tyr Lys Gly Ser Leu Thr 395 390 339 Ile Phe Ala Leu Asn Val Gly Asp Gly Asp Val Thr Leu Lys Ile Gly 405 410 342 Gln Tyr Ser Gly Lys Lys Ile Tyr Ser Tyr Ile Leu Thr Pro Glu Gly 420 425 345 Gly Gln Leu Thr Ser Gln Lys Val Leu Leu Asn Gly Lys Glu Leu Asn 440 435 348 Leu Val Ser Asp Gln Leu Pro Glu Leu Asn Ala Asp Glu Ser Lys Thr 460 450 455 351 Ser Phe Thr Leu Ser Pro Lys Thr Phe Gly Phe Phe Val Val Ser Asp 470 475 354 Ala Asn Val Glu Ala Cys Lys Lys 355 485 498 <210> SEQ ID NO: 5 499 <211> LENGTH: 488 500 <212> TYPE: PRT 501 <213> ORGANISM: Leech 503 <400> SEQUENCE: 5 504 Lys Glu Ile Ala Val Thr Ile Asp Asp Lys Asn Val Ile Ala Ser Ala 5 10 507 Ser Glu Ser Phe His Gly Val Ala Phe Asp Ala Ser Leu Phe Ser Pro 2:0 25 510 Lys Gly Leu Trp Ser Phe Val Asp Ile Thr Ser Pro Lys Leu Phe Lys 513 Leu Leu Glu Gly Leu Ser Pro Gly Tyr Phe Arg Val Gly Gly Thr Phe

Jan Haraber



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514		50					55					60				
	Ala		Ara	Leu	Phe	Phe		Leu	Asp	Glu	Asn	Asn	Lvs	Trp	Lys	Asp
517	65		,			70					75		1	•	•	80
519	Tyr	Trp	Ala	Phe	Lys	Asp	Lys	Thr	Pro	Glu	Thr	Ala	Thr	Ile	Thr	Arg
520	1	_			85	•	1			90					95	,
	Ara	Trp	Leu	Phe	Ara	Lvs	Gln	Asn	Asn	Leu	Lvs	Lvs	Glu	Thr	Phe	Asp
523	,	1	-	100	,	1 -		-	105		1	1		110		
	Asn	Leu	Val	Lvs	Leu	Thr	Lvs	Glv	Ser	Lvs	Met	Arq	Leu	Leu	Phe	Asp
526			115	1			1	120		1		,	125			-
	Leu	Asn		Glu	Va1	Arg	Thr	Glv	Tyr	Glu	Ile	Glv	Lvs	Lys	Met	Thr
529		130				,	135	1	1			140	1	4		
531	Ser	Thr	Trp	Asp	Ser	Ser	Glu	Ala	Glu	Lys	Leu	Phe	Lys	Tyr	Cys	Val
	145		_	•		150				-	155		•	1		160
534	Ser	Lys	Gly	Tyr	Gly	Asp	Asn	Ile	Asp	Trp	Glu	Leu	Gly	Asn	Gly	Pro
535		4	1	1	165	1			-	170					175	
537	Asp	His	Thr	Ser	Ala	His	Asn	Leu	Thr	Glu	Lys	Gln	Val	Gly	Glu	Asp
538	-			180					185		•			190		•
540	Phe	Lys	Ala	Leu	His	Lys	Val	Leu	Glu	Lys	Tyr	Pro	Thr	Leu	Asn	Lys
541		1	195			1		200		-	•		205			•
543	Gly	Ser	Leu	Val	Gly	Pro	Asp	Val	Gly	Trp	Met	Gly	Val	Ser	Tyr	Val
544	•	210			_		215		•	-		220			•	
546	Lvs	Gly	Leu	Ala	Asp	Glu	Ala	Gly	Asp	His	Val	Thr	Ala	Phe	Thr	Leu
	225	•			•	230		_	•		235					240
549	His	Gln	Tyr	Tyr	Phe	Asp	Gly	Asn	Thr	Ser	Asp	Val	Ser	Ile	Tyr	Leu
550			•	•	245	•	-			250	•				255	
552	Asp	Ala	Thr	Tyr	Phe	Lys	Lys	Leu	Gln	Gln	Leu	Phe	Asp	Lys	Val	Lys
553	-			260		-	-		265					270		
555	Asp	Val	Leu	Lys	Asp	Ser	Pro	His	Lys	Asp	Lys	Pro	Leu	Trp	Leu	Gly
556	_		275	_	_			280	_	_	_		285			
558	Glu	Thr	Ser	Ser	Gly	Tyr	Asn	Ser	Gly	Thr	Glu	Asp	Val	Ser	Asp	Arg
559		290			_	_	295					300				
561	Tyr	Val	Ser	Gly	Phe	Leu	Thr	Leu	Asp	Lys	Leu	Gly	Leu	Ser	Ala	Ala
562	305					310					315					320
564	Asn	Asn	Val	Lys	Val	Val	Ile	Arg	Gln	Thr	Ile	Tyr	Ser	Gly	Tyr	Tyr
565					325					330					335	
567	Gly	Pro	Leu	Asp	Lys	Asn	Thr	Leu	$\operatorname{Glu}$	Pro	Asn	Pro	Asp	Tyr	Trp	Leu
568				340					345					350		
570	Met	His	Val	His	Asn	Ser	Leu	Val	Gly	Asn	Thr	Val	Phe	Lys	Val	Asp
571			355					360					365			
573	Val	Ser	Asp	Pro	Thr	Asn	Lys	Ala	Arg	Val	Tyr	Ala	Gln	Cys	Thr	Lys
574		370					375					380				
576	Thr	Asn	Ser	Lys	His	Thr	Gln	Ser	Arg	Tyr	Tyr	Lys	Gly	Ser	Leu	Thr
577	385					390					395					400
579	Ile	Phe	Ala	Leu	Asn	Val	Gly	Asp	Glu	Asp	Val	Thr	Leu	Lys	Ile	Gly
580					405					410					415	
	Gln	Tyr	Ser	Gly	Lys	Lys	Ile	Tyr		Tyr	Ile	Leu	Thr	Pro	Glu	Gly
583				420					425					430		
	Gly	Gln		Thr	Ser	Gln	Lys	Val	Leu	Leu	Asn	Gly	_	Glu	Leu	Asn
586			435					440					445			

RAW SEQUENCE LISTING

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Input Set : A:\merck 2332 seq list

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E--> 588 Let Xaa Ser Asp Gln Leu Pro Gln Leu Asn Ala Asp Glu Ser Lys Thr

589 450 455 460

 $591~{\rm Ser}$  Phe Thr Leu Ser Pro Lys Thr Phe Gly Phe Phe Val Val Ser Asp

592 465 470 475

594 Ala Asn Val Glu Ala Cys Lys Lys

595 485

VARIABLE LOCATION SUMMARY

PATENT APPLICATION: US/10/009,500

DATE: 06/06/2002 TIME: 14:28:18

Input Set : A:\merck 2332 seq list

Output Set: N:\CRF3\06062002\J009500.raw

Use of n's or Xaa's (NEW RULES):

Use of n's and/or Xaa's have been detected in the Sequence Listing. Use of  $\langle 220 \rangle$  to  $\langle 223 \rangle$  is MANDATORY if n's or Xaa's are present. in  $\langle 220 \rangle$  to  $\langle 223 \rangle$  section, please explain location of n or Xaa, and which residue n or Xaa represents.

Seq#:2; Xaa Pos. 223,231,236,249

Seq#:3; Xaa Pos. 223,231,236,249

Seq#:4; Xaa Pos. 450

Seq#:5; Xaa Pos. 450



## VERIFICATION SUMMARY

PATENT APPLICATION: US/10/009,500

DATE: 06/06/2002 TIME: 14:28:18

Input Set : A:\merck 2332 seq list

Output Set: N:\CRF3\06062002\J009500.raw

L:11 M:270 C: Current Application Number differs, Replaced Application Number L:12 M:271 C: Current Filing Date differs, Replaced Current Filing Date

L:186 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:2 after pos.:672

 $L:190\ M:341\ W:$  (46) "n" or "Xaa" used, for SEQ ID#:2 after pos.:720 L:194 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:2 after pos.:768

L:303 M:340 E: (46) "n" or "Xaa" used: Feature required, for SEQ ID#:3

M:340 Repeated in SeqNo=3

 $L:486\ M:341\ W:$  (46) "n" or "Xaa" used, for SEQ ID#:4 after pos.:1392

L:588 M:340 E: (46) "n" or "Xaa" used: Feature required, for SEQ ID#:5